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### NOTICE OF ALLOWANCE AND FEE(S) DUE

49002 7590 9329/2012 JOYCE VON NATZMER PEQUIGNOT + MYERS LLC 200 Madison Avenue Suite 1901 New York, NY 10016 EXAMINER

QIAN, CELINE X

APT INST

PAGED NUMBED

1636 DATE MAILED: 03/29/2012

APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
10/595,495	04/24/2006	Nicolas Mermod	3024-119	1575

APPLN, TYPE	SMALL ENTITY	ISSUE FEE DUE	PUBLICATION FEE DUE	PREV. PAID ISSUE FEE	TOTAL FEE(S) DUE	DATE DUE
nonprovisional	NO	\$1740	\$300	\$0	\$2040	06/29/2012

THE APPLICATION IDENTIFIED ABOVE HAS BEEN EXAMINED AND IS ALLOWED FOR ISSUANCE AS A PATENT. PROSECUTION ON THE MERITS IS CLOSED. THIS NOTICE OF ALLOWANCE IS NOT A GRANT OF PATENT RIGHTS. THIS APPLICATION IS SUBJECT TO WITHDRAWAL FROM ISSUE AT THE INITIATIVE OF THE OFFICE OR UPON PETITION BY THE APPLICANT. SEE 37 CFR 1.313 AND MPEP 1308.

THE ISSUE FEE AND PUBLICATION FEE (IF REQUIRED) MUST BE PAID WITHIN THREE MONTHS FROM THE MAILING DATE OF THIS NOTICE OR THIS APPLICATION SHALL BE REGARDED AS ABANDONED. THIS STATUTORY PERIOD CANNOT BE EXTENDED. SEE 35 U.S.C. 1SI. THE ISSUE FEE DUE INDICATED ABOVE DOES NOT REFLECT A CREDIT FOR ANY PREVIOUSLY PAID ISSUE FEE IN THIS APPLICATION. IF AN ISSUE FEE HAS PREVIOUSLY BEEN PAID IN THIS APPLICATION (AS SHOWN ABOVE), THE RETURN OF PART B OF THIS FORM WILL BE CONSIDERED A REQUEST TO REAPPLY THE PREVIOUSLY PAID ISSUE FEE TOWARD THE ISSUE FEE NOW DUE.

#### HOW TO REPLY TO THIS NOTICE:

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If the SMALL ENTITY is shown as NO:

A. Pay TOTAL FEE(S) DUE shown above, or

B. If applicant claimed SMALL ENTITY status before, or is now claiming SMALL ENTITY status, check box 5a on Part B - Fee(s) Transmittal and pay the PUBLICATION FEE (if required) and 1/2 the ISSUE FEE shown above.

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III. All communications regarding this application must give the application number. Please direct all communications prior to issuance to Mail Stop ISSUE FEE unless advised to the contrary.

IMPORTANT REMINDER: Utility patents issuing on applications filed on or after Dec. 12, 1980 may require payment of maintenance fees. It is patentee's responsibility to ensure timely payment of maintenance fees when due.

#### PART B - FEE(S) TRANSMITTAL

Complete and send this form, together with applicable fee(s), to: Mail Mail Stop ISSUE FEE Commissioner for Patents P.O. Box 1450 Alexandria, Virginia 22313-1450

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03/29/2012

JOYCE VON NATZMER PEOUIGNOT + MYERS LLC 200 Madison Avenue Suite 1901

New York, NY 10016

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(Depositor's nam (Signate (Dat

APPLICATION NO FILING DATE FIRST NAMED INVENTOR ATTORNEY DOCKET NO CONFIRMATION NO 10/595 495 04/24/2006 Nicolas Mermod 3024-119 1575 TITLE OF INVENTION: HIGH EFFICIENCY GENE TRANSFER AND EXPRESSION IN MAMMALIAN CELLS BY A MULTIPLE TRANSFECTION

PROCEDURE FO MAR SEQUENCES

APPLN. TYPE	SMALL ENTITY	ISSUE FEE DUE	PUBLICATION FEE DUE	PREV. PAID ISSUE FEE	TOTAL FEE(S) DUE	DATE DUE
nonprovisional	NO	\$1740	\$300	\$0	\$2040	06/29/2012
EXAM	IINER	ART UNIT	CLASS-SUBCLASS			
QIAN, CI	ELINE X	1636	536-024100	•		
I. Change of correspondence address or indication of "Fee Address" (37 CFR 1.363).  CFR 1.363).  Change of correspondence address (or Change of Correspondence Address form PTOSB/122) attached.  "Fee Address" indication (or "Fee Address" Indication form PTOSB/147; Rev U3-U2 or more recent) attached. Use of a Customer Number is required.			or agents OR, alternativ (2) the name of a single registered attorney or a	3 registered patent attorney, ely, e firm (having as a memb igent) and the names of up meys or agents. If no nam	era 2	

Number is required.	inter, no mane with the prince.
3. ASSIGNEE NAME AND RESIDENCE DATA TO BE PRINTED ON T	THE PATENT (print or type)
PLEASE NOTE: Unless an assignee is identified below, no assignee recordation as set forth in 37 CFR 3.11. Completion of this form is NO	data will appear on the patent. If an assignee is identified below, the document has been filed for $\Gamma$ a substitute for filing an assignment.
(A) NAME OF ASSIGNEE	(B) RESIDENCE: (CITY and STATE OR COUNTRY)
Please check the appropriate assignee category or categories (will not be pr	inted on the patent): 🔲 Individual 🚨 Corporation or other private group entity 🚨 Government
4. The fell and a fee(a) are submitted.	Description of the control of the co
4a. The following fee(s) are submitted: 4b □ Issue Fee	Payment of Fee(s): (Please first reapply any previously paid issue fee shown above)     A check is enclosed.
Publication Fee (No small entity discount permitted)	Payment by credit card. Form PTO-2038 is attached.
Advance Order - # of Copies	The Director is hereby authorized to charge the required fee(s), any deficiency, or credit any overpayment, to Deposit Account Number(enclose an extra copy of this form).
5. Change in Entity Status (from status indicated above)	
a. Applicant claims SMALL ENTITY status. See 37 CFR 1.27.	☐ b. Applicant is no longer claiming SMALL ENTITY status. See 37 CFR 1.27(g)(2).
NOTE: The Issue Fee and Publication Fee (if required) will not be accepted interest as shown by the records of the United States Patent and Trademark	d from anyone other than the applicant; a registered attorney or agent; or the assignee or other party in Office.

Authorized Signature Date Typed or printed name Registration No. This collection of information is required by 37 CFR 1.311. The information is required to obtain or retain a benefit by the public which is to file (and by the USPTO to process) an application. Confidentiality is governed by 35 U.S.C. 122 and 37 CFR 1.14. This collection is estimated to take 12 minutes to complete, including gathering, preparing, and submitting the completed application form to the USPTO. Time will vary depending upon the individual case. Any comments on the amount of time you require to complete this form and/or suggestions for rectucing this burden, should be sent to the Chief Information Officer. U.S. Patest and Trademark Officer. U.S. Department of Commerce, P.O. Box 1450, Alexandria, Virginia 2231-450. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Commissioner for Patents, P.O. Box 1450, Alexandria, Virginia 2231-450.

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# UNITED STATES PATENT AND TRADEMARK OFFICE

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 Nicolas Mermod
 3/024-119
 1575

JOYCE VON NATZMER
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200 Madison Avenue
Suite 1901

New York, NY 10016

QIAN, CELINE X

ART UNIT PAPER NUMBER

1636

DATE MAILED: 03/29/2012

# Determination of Patent Term Adjustment under 35 U.S.C. 154 (b)

(application filed on or after May 29, 2000)

The Patent Term Adjustment to date is 169 day(s). If the issue fee is paid on the date that is three months after the mailing date of this notice and the patent issues on the Tuesday before the date that is 28 weeks (six and a half months) after the mailing date of this notice, the Patent Term Adjustment will be 169 day(s).

If a Continued Prosecution Application (CPA) was filed in the above-identified application, the filing date that determines Patent Term Adjustment is the filing date of the most recent CPA.

Applicant will be able to obtain more detailed information by accessing the Patent Application Information Retrieval (PAIR) WEB site (http://pair.uspto.gov).

Any questions regarding the Patent Term Extension or Adjustment determination should be directed to the Office of Patent Legal Administration at (571)-272-7702. Questions relating to issue and publication fee payments should be directed to the Customer Service Center of the Office of Patent Publication at 1-(888)-786-0101 or (571)-272-4200.

## Privacy Act Statement

The Privacy Act of 1974 (P.L. 93-579) requires that you be given certain information in connection with your submission of the attached form related to a patent application or patent. Accordingly, pursuant to the requirements of the Act, please be advised that: (1) the general authority for the collection of this information is 35 U.S.C. 2(b)(2); (2) furnishing of the information solicited is voluntary; and (3) the principal purpose for which the information is used by the U.S. Patent and Trademark Office is to process and/or examine your submission related to a patent application or patent. If you do not furnish the requested information, the U.S. Patent and Trademark Office may not be able to process and/or examine your submission, which may result in termination of proceedings or abandonment of the application or expiration of the patent.

The information provided by you in this form will be subject to the following routine uses:

- The information on this form will be treated confidentially to the extent allowed under the Freedom
  of Information Act (5 U.S.C. 552) and the Privacy Act (5 U.S.C 552a). Records from this system of
  records may be disclosed to the Department of Justice to determine whether disclosure of these
  records is required by the Freedom of Information Act.
- A record from this system of records may be disclosed, as a routine use, in the course of presenting evidence to a court, magistrate, or administrative tribunal, including disclosures to opposing counsel in the course of settlement negotiations.
- 3. A record in this system of records may be disclosed, as a routine use, to a Member of Congress submitting a request involving an individual, to whom the record pertains, when the individual has requested assistance from the Member with respect to the subject matter of the record.
- 4. A record in this system of records may be disclosed, as a routine use, to a contractor of the Agency having need for the information in order to perform a contract. Recipients of information shall be required to comply with the requirements of the Privacy Act of 1974, as amended, pursuant to 5 U.S.C. 552a(m).
- 5. A record related to an International Application filed under the Patent Cooperation Treaty in this system of records may be disclosed, as a routine use, to the International Bureau of the World Intellectual Property Organization, pursuant to the Patent Cooperation Treaty.
- A record in this system of records may be disclosed, as a routine use, to another federal agency for purposes of National Security review (35 U.S.C. 181) and for review pursuant to the Atomic Energy Act (42 U.S.C. 218(c)).
- 7. A record from this system of records may be disclosed, as a routine use, to the Administrator, General Services, or his/her designee, during an inspection of records conducted by GSA as part of that agency's responsibility to recommend improvements in records management practices and programs, under authority of 44 U.S.C. 2004 and 2006. Such disclosure shall be made in accordance with the GSA regulations governing inspection of records for this purpose, and any other relevant (i.e., GSA or Commerce) directive. Such disclosure shall not be used to make determinations about individuals
- 8. A record from this system of records may be disclosed, as a routine use, to the public after either publication of the application pursuant to 35 U.S.C. 122(b) or issuance of a patent pursuant to 35 U.S.C. 151. Further, a record may be disclosed, subject to the limitations of 37 CFR 1.14, as a routine use, to the public if the record was filed in an application which became abandoned or in which the proceedings were terminated and which application is referenced by either a published application, an application open to public inspection or an issued patent.
- A record from this system of records may be disclosed, as a routine use, to a Federal, State, or local law enforcement agency, if the USPTO becomes aware of a violation or potential violation of law or regulation.

## Application No. Applicant(s) 10/595.495 MERMOD ET AL. Notice of Allowability Examiner Art Unit CELINE CIAN 1636 -- The MAILING DATE of this communication appears on the cover sheet with the correspondence address--All claims being allowable, PROSECUTION ON THE MERITS IS (OR REMAINS) CLOSED in this application. If not included herewith (or previously mailed), a Notice of Allowance (PTQL-85) or other appropriate communication will be mailed in due course. THIS NOTICE OF ALLOWABILITY IS NOT A GRANT OF PATENT RIGHTS. This application is subject to withdrawal from issue at the initiative of the Office or upon petition by the applicant. See 37 CFB 1.313 and MPEP 1308. This communication is responsive to the response filed on 2/9/2012. 2. An election was made by the applicant in response to a restriction requirement set forth during the interview on the restriction requirement and election have been incorporated into this action. 3. X The allowed claim(s) is/are 48.49.51.55.65.67.68,70-72.74-91.103.105-107.110-115.117-119 and 122-126. Acknowledgment is made of a claim for foreign priority under 35 U.S.C. § 119(a)-(d) or (f). b) Some\* c) None of the: Certified copies of the priority documents have been received. 2. Certified copies of the priority documents have been received in Application No. 3. Copies of the certified copies of the priority documents have been received in this national stage application from the International Bureau (PCT Rule 17.2(a)). \* Certified copies not received: Applicant has THREE MONTHS FROM THE "MAILING DATE" of this communication to file a reply complying with the requirements noted below. Failure to timely comply will result in ABANDONMENT of this application. THIS THREE-MONTH PERIOD IS NOT EXTENDABLE. 5. A SUBSTITUTE OATH OR DECLARATION must be submitted. Note the attached EXAMINER'S AMENDMENT or NOTICE OF INFORMAL PATENT APPLICATION (PTO-152) which gives reason(s) why the oath or declaration is deficient. 6. CORRECTED DRAWINGS ( as "replacement sheets") must be submitted. (a) I including changes required by the Notice of Draftsperson's Patent Drawing Review (PTO-948) attached 1) Thereto or 2) to Paper No./Mail Date (b) including changes required by the attached Examiner's Amendment / Comment or in the Office action of Paper No./Mail Date Identifying indicia such as the application number (see 37 CFR 1.84(c)) should be written on the drawings in the front (not the back) of each sheet. Replacement sheet(s) should be labeled as such in the header according to 37 CFR 1.121(d). 7. \$\scale\$ DEPOSIT OF and/or INFORMATION about the deposit of BIOLOGICAL MATERIAL must be submitted. Note the attached Examiner's comment regarding REQUIREMENT FOR THE DEPOSIT OF BIOLOGICAL MATERIAL. Attachment(s) 1. Notice of References Cited (PTO-892) 5. Notice of Informal Patent Application

2. Notice of Draftperson's Patent Drawing Review (PTO-948)

4. T Examiner's Comment Regarding Requirement for Deposit

Information Disclosure Statements (PTO/SB/08).

Paper No./Mail Date

of Biological Material

 Interview Summary (PTO-413), Paper No./Mail Date 0312.

9. ☐ Other .

 ☐ Examiner's Amendment/Comment

8. T Examiner's Statement of Reasons for Allowance

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### EXAMINER'S AMENDMENT

An examiner's amendment to the record appears below. Should the changes and/or additions be unacceptable to applicant, an amendment may be filed as provided by 37 CFR 1.312. To ensure consideration of such an amendment, it MUST be submitted no later than the payment of the issue fee.

Authorization for this examiner's amendment was given in a telephone interview with Applicant's representative Ms. Joyce v. Natzmer on 3/23/2012.

The application has been amended as follows:

Claims 15-17, 23, 24, 26, 28, 34, 42-45, 62-64, 69, 92, 93, 97-100 and 108 are canceled.

Claim 48. A method for transfecting a eukaryotic host cell, said method comprising a) introducing into said eukaryotic host cell at least one purified DNA sequence comprising at least one DNA sequence of interest and/or-at least one first purified and isolated DNA sequence according to claim 65, b) subjecting said transfected eukaryotic host cell to at least one additional transfection step with at least one second purified DNA sequence of interest and/or with at least one second purified and isolated DNA sequence comprising a MAR nucleotide sequence or other chromatin modifying elements, and c) selecting said transfected eukaryotic host cell.

Claim 77. The vector of claim 74 75, wherein said vector further comprises one or more regulatory sequences.

Claim 85. A transfected eukaryotic host cell transfected with at least one vector comprising at least one purified DNA sequence comprising:

a) at least one bent DNA element comprising at least 33% of the dinucleotide TA and/or at least 33% of the dinucleotide AT on a stretch of 100 contiguous base pairs. Application/Control Number: 10/595,495

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b) at least one binding site for a DNA binding protein, wherein said purified DNA sequence is a MAR nucleotide sequence with sequence ID NO: 25, or a sequence complementary to sequence ID NO: 25, or a sequence having at least 90% identity with said SEQ ID NO: 25.

Claim 124. A purified and isolated DNA sequence, wherein said DNA sequence comprises a) at least one bent DNA element, b) and at least one binding site for a DNA binging binding protein, and that it wherein said DNA sequence is a MAR nucleotide with SEQ ID NO: 25, or a sequence complementary thereof, or a sequence having at least 90% identity with said sequence.

Claim 125. The vector of claim 82 78, wherein said first DNA sequence and said second DNA sequence are located on a sequence distinct from the one containing a promoter and said gene of interest.

Claim 126. The host cell according to claim 85, wherein the host cell is a high recombinant protein produced producing cell with a production rate of at least 10pg per cell per day.

Title: High efficiency gene transfer and expression in mammalian cells by multiple transfection procedure of fo MAR sequences

Any inquiry concerning this communication or earlier communications from the examiner should be directed to CELINE QIAN whose telephone number is (571)272-0777. The examiner can normally be reached on 9:30-5:30 M-F.

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If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Ardin Marschel can be reached on 571-272-2911. The fax phone number for the organization where this application or proceeding is assigned is 571-273-8300.

Information regarding the status of an application may be obtained from the Patent Application Information Retrieval (PAIR) system. Status information for published applications may be obtained from either Private PAIR or Public PAIR. Status information for unpublished applications is available through Private PAIR only. For more information about the PAIR system, see http://pair-direct.uspto.gov. Should you have questions on access to the Private PAIR system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free). If you would like assistance from a USPTO Customer Service Representative or access to the automated information system, call 800-786-9199 (IN USA OR CANADA) or 571-272-1000.

/Celine X Qian / Primary Examiner, Art Unit 1636